I hereby certify that this correspondence is being deposited with the US Postal Service with sufficient postage as the class mail in an envelope addressed to the Mail Stop DD, Commissioner for Patents, P. O. Box 1450, Alexandria, VA 22313-450, on the date shown below.

Date: September 4, 2003

By: September 4, 2003

PATENT Docket No. GC541-4-C1

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re A Bott et	pplication of al.)	
•	10/629,976) Group Art Unit: Not Assigne)	ď
Serial No.: Unassigned) Examiner: Not Assigned	
Filed:	Herewith)	
For:	SYNTHESIS AND USE OF GLYCODENDRIMER REAGENTS))	

Information Disclosure Statement

Mail Stop DD Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Sir:

Applicants submit herewith patents, publications or other information (listed on the attached Form PTO-1449 and attached thereto) of which they are aware, that they believe may be material to the examination of this application and in respect of which there may be a duty to disclose in accordance with 37 CFR §1.56.

This Information Disclosure Statement:

(a) accompanies the new patent application submitted herewith. 37 CFR §1.97(a).
(b) is filed within three months after the filing date of the application or within three months after the date of entry into the national stage of a PCT application as set forth in 37 CFR §1.491.
(c) \square as far as is known to the undersigned, is filed before the mailing date of a first Office Action on the merits.

(d) is filed after the first Office Action and more than three months after the application filing date or PCT national stage date of entry filing but, as far

	rejection either specific Depos Inform	nown to the undersigned, prior to the mailing date of either a final on or a notice of allowance, whichever occurs first, and is accompanied by the fee (\$180.00) set forth in 37 CFR §1.17(p) or a certification as ed in 37 CFR §1.97(e), as checked below. Authorization to charge sit Account No. 07-1048 in the amount of \$180.00 to cover the cost of this ation Disclosure Statement is provided in the Transmittal Letter submitted ith in duplicate.
	allowa Transr No. 07 as spe consid	is filed after the mailing date of either a final rejection or a notice of nce, whichever occurred first, and is accompanied by authorization (in the mittal Letter submitted herewith in duplicate) to charge Deposit Account 7-1048 the fee (\$180.00) set forth in 37 CFR §1.17(I)(1) and a certification ecified in 37 CFR §1.97(e), as checked below. This document is to be dered as a petition requesting consideration of the Supplemental nation Disclosure Statement.
[If either of b	oxes (c	d) or (e) is checked above, the following "certification" under 37 CFR
§1.97(e) may	need t	o be completed.] The undersigned certifies that:
	counte	Each item of information contained in the Information Disclosure nent was cited in a communication mailed from a foreign patent office in a expart foreign application not more than three months prior to the filing of formation Disclosure Statement.
	foreigr reasor	No item of information contained in this Information Disclosure Statement ted in a communication mailed from a foreign patent office in a counterpart application or, to the knowledge of the undersigned after making hable inquiry, was known to any individual designated in 37 CFR §1.56(c) than three months prior to the filing of this Information Disclosure nent.
Those patent	(s) or pu	ublication(s) which are marked with an asterisk (*) on the attached Form
PTO-1449 are	e not su	pplied because they were previously cited by or submitted to the Office in
a prior applica	ation, Se	erial No. 09/824,827 filed April 2, 2001.
A concise exp	lanatio	n of relevance of the items listed on PTO-1449 is:
	\boxtimes	not given
		given for each listed item
		given for only non-English language listed item(s)
	_	in the form of an English language copy of a Search Report from a patent office, issued in a counterpart application, which refers to the nt portions of the references.

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While the information and references disclosed in this Information Disclosure Statement may be "material" pursuant to 37 CFR §1.56, it is not intended to constitute an admission that any patent, publication or other information referred to therein is "prior art" for this invention unless specifically designated as such.

In accordance with 37 CFR §1.97(b), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 CFR §1.56(a) exists. It is submitted that the Information Disclosure Statement is in compliance with 37 CFR §1.98 and MPEP §609 and the Examiner is respectfully requested to consider the listed references.

Respectfully submitted,

Date: September 4, 2003

H. Thomas Anderton, Jr. Registration No. 40,895

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Tel: 650 846-7544 Fax: 650 845-6504

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(PTO-1449)

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SERIAL NO.

10/629,976

APPLICANT

Bott et al. FILING DATE

July 30, 2003

GROUP ART UNIT

Unassigned

REFERENCE DESIGNATION	U.S. PATENT DOCUMENTS

EXAM'R INITIAL		DOCUMENT NUMBER	DATE	NAME	Class	Subclass	Filing Date If Appropriate
	A1	*5,403,737	04/04/95	Abrahmsen et al.			
	A2	*5,629,173	05/13/97	Abrahmsen et al.			
	A3	*5,316,935	05/31/94	Arnold et al.			
	A4	*5,208,158	05/04/93	Bech et al.			
	A5	*5,244,791	09/14/93	Estell			
	A6	*5,316,941	05/31/94	Estell et al.			
	A7	*5,955,340	02/21/99	Bott			
	A8	*5,340,735	08/23/94	Christianson et al.			

FOREIGN PATENT DOCUMENTS

EXAM'R INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	Subclass	TRANSLAT'N
	B1	*EP 3 328 229 A1	08/16/89	EP		,	
	B2	*WO 00/01712	01/13/00	PCT			
	B3	*WO 91/16423	04/18/91	PCT			
	B4	*WO 96/27671	02/27/96	PCT			
	B5	*WO 97/37007	10/09/97	PCT			
	B6	*WO 98/23732	06/04/98	PCT			
	B7	*WO 99/20723	04/29/99	PCT			
	B8	*WO 99/37323	07/29/99	PCT			
	B9	*WO 99/37324	07/29/99	PCT			

OTHER DOCUMENTS (including Author, Title, Date, Pertinent Pages, Etc.)

C1	*Bech et al., "Chemical Modifications of a Cysteinyl Residue Introduced in the Binding Site of Carboxypeptidase Y by Site-Directed Mutagenesis," Carlsberg Res. Commun., 53:381-393 (1988)
C2	*Bech et al., "Significance of Hydrophobic S ₄ -P ₄ Interactions in Subtilisin 309 from <i>Bacillus Ientus</i> ," Biochemistry, 32:2847-2852 (1993)
СЗ	*Berglund et al., "Altering the Specificity of Subtilisin B. Lentus by Combining Site-Directed Mutagenesis and Chemical Modification," <u>Bioorganic & Mechanical Chemistry Letters</u> , 6:2507-2512 (1996)
 C4	*Berglund et al., "Chemical Modification of Cysteine Mutants of Subtilisin <i>Bacillus Lentus</i> Can Create Better Catalysts Than The Wild-Type Enzyme," J. Am. Chem. Soc., 119:5265-5266 (1997)

EXAMINER

DATE CONSIDERED

EXAMINER: Initial if reference considered, whether or not citation is in c nformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant(s).

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APPLICANT Bott et al. FILING DATE

GROUP ART UNIT

July 30, 2003 Unassigned

	OTHER DOCUMENTS (including Author, Title, Date, Pertinent Pages, Etc.)
C5	*Betzel et al., "Crystal Structure of the Alkaline Proteinase Savinase TM from <i>Bacillus lentus</i> at 1 4 Å Resolution," <u>J. Mol. Biol.</u> , 223:427-445(1992)
C6	*Bonneau et al., "Alteration of the Specificity of Subtilisin BPN' by Site-Directed Mutagenesis in its S ₁ and S ₁ ' Binding Sites," <u>J. Am. Chem. Soc.</u> , 113:1026-30 (1991)
C7	*Brocklehurst, "Specific Covalent Modification of Thiols: Applications in the Study of Enzymes and Other Biomolecules," Int. J. Biochem., 10:259-274 (1979)
C8	*Bruice et al., "Novel Alkyl Alkanethiolsulfonate Sulfhydryl Reagents. Modification of Derivatives of L-Cysteine," <u>Journal of Protein Chemistry</u> , 1:47-58 (1982)
C9	*Chen et al., "Probing the S-1' Subsite Selectivity of an Industrial Alkaline Protease in Anhydrous t-Butanol," <u>Bioorganic & Medicinal Chemistry Letters</u> , 3(4):727-33 (1993)
C10	*Davies et al., "A Semisynthetic Metalloenzyme Based on a Protein Cavity That Catalyzes the Enantiosleective Hydrolysis of Ester and Amide Substrates," <u>J. Am. Chem. Soc.</u> , 119:11643-11652 (1997)
C11	*Davis, B.G., et al., "Altering the specificity of subtilisin Bacillus lentus through the introduction of positive charge at single amino acid sites," <u>Bioorganic and Medicinal Chemistry</u> , (1999 Nov.) 7 (11) 2303-11, XPO000892841
C12	*Davis, B.G., et al., "Controlled site selective protein glycosylation for precise glycan structure catalytic activity relationships," Biorganic & Medicinal Chemistry, Vol. 8, 2000, pp. 1527-1535
C13	*Davis, B.G., et al., "Glycomethanethiosulfonates: powerful reagents for protein glycosylation," Tetrahedron: Asymmetry, NL, Elsevier Science Publishers, Amsterdam, Vol 11, No. 1, January 2000 (2000-01), pp. 245-262
C14	*Davis, B.G., et al., "The controlled introduction of multiple negative charge at single amino acid sites in subtilisin bacillus lentus," <u>Bioorganic and Medicinal Chemistry</u> , (1999 Nov.) 7 (11) 2293-301, XPO000892840
C15	Davis, B.G., et al., "Glycosyldisulfides: a new class of solution and solid phase glycosyl donors," Chem. Commun, 2001, pp.189-190
C16	*Davis, Benjamin G, et al., "Controlled Site Selective Glycosylation of Proteins by a Combined Site Directed Mutagenesis and Chemical Modification Approach," <u>J. Org. Chem.</u> , Vol. 63, January 12, 1998 (1998-01-12), pp. 9614-9615
C17	*Davis, Benjamin G, et al., "The Controlled Glycosylation of a Protein with a Bivalent Glycan: Towards a New Class of Glycoconjuates, Glycodendriproteins," Chem. Commun, 2001, pp. 351-352
C18	*DeSantis et al., "Chemical Modifications at a Single Site Can Induce Significant Shifts in the pH Profiles of a Serine Protease," <u>J. Am Chem. Soc.</u> , 120:8582-8586 (1998)
C19	*Desantis, G., et al, "Probing the altered specificity and catalytic properties of mutant subtilisin chemically modified at position S156C and S166C in the S1 pocket," Bioorganic and Medicinal Chemistry, (1997) 7/7 (1381-1387), XP0000892843

EXAMINER DATE CONSIDERED

EXAMINER: Initial if reference c nsidered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy f this form with next communication to Applicant(s).

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(PTO-1449)

GROUP ART UNIT FILING DATE July 30, 2003 Unassigned

	OTHER DOCUMENTS (including Author, Title, Date, Pertinent Pages, Etc.)
C20	*DeSantis, G., et al., "Site-Directed Mutagenesis Combined with Chemical Modification As a Strategy for Altering the Specificity of the S1 and S1' Pockets of Subtilisin Bacillus Lentus," <i>Biochemistry</i> (1998) 37 (17) 5968-73
C21	*Dickman, M., et al., "Chemically modified mutants of subtilisin bacillus lentus catalyze transesterification reactions better than wild type," <u>Tetrahedron Asymmetry</u> , (11. Dec. 1998) 9/23 4099-4102, XPO000901276.
C22	*Gron et al., "A Highly Active and Oxidation-Resistant Subtilisin-Like Enzyme Produced by a Combination of Site-Directed Mutagenesis and Chemical Modification," <u>Eur. J. Biochem.</u> , 194:897-901 (1990)
C23	*Kaiser, "Catalytic Activity of Enzymes Altered at Their Active Sites," Agnew. Chem. Int. Ed. Engl., 27-913-922 (1988)
C24	*Kawase et al., "Effect of Chemical Modification of Tyrosine Residues on Activities of Bacterial Lipase," <u>Journal of Fermentation and Bioengineering</u> , 72:317-319 (1991)
C25	*Kenyon et al., "Novel Sulfhydryl Reagents," Methods Enzymol., 47:407-430 (1977)
C26	*Kluger et al., "Amino Group Reactions of the Sulfhydryl Reagent Methyl Methanesulfonothioate. Inactivation of D-3-hydroxybutyrate Dehydrogenase and Reaction with Amines in Water," <u>Can. J. Biochem.</u> , 58:629-632 (1980)
C27	*Lloyd, R.C. et al., "Site Selective Glycosilation of Subtilisin Bacillus Lentus Causes Dramatic Increase in Esterase Activity," <u>Biorganic & Medicinal Chemistry</u> , Vol. 8, 2000, pp. 1537-1544
C28	*Lo, Bryan, et al., "Replacement of Ala-166 with Cysteine in the High Affinity Rabbit Sodium Glucose Transporter Alters Transport Kinetics and Allows Methanethiosulfonate Ethylamine to Inhibit Transporter Function," The Journal of Biological Chemistry, 273:2 903-909 (1998)
C29	*Neet, K.E. and Koshland, D.E., "The Conversion of Serine at the Active Site of Subtilisin to Cysteine: A 'Chemical Mutation," Proc. Nat. Acad. Sci. USA, 56(5):1606-1611.
C30	*Nishimura et al., "Reversible Modification of the Sulfhydryl Groups of <i>Escherichia coli</i> Succinic Thiokinase with Methanethiolating Reagents, 5,5'-Dithio-bis(2-Nitrobenzoic Acid), p-Hydroxymercuribenzoate, and Ethylmercurithiosalicylate," <u>Archives of Biochemistry and Biophysics</u> , 170:461-467 (1975)
C31	*Paulson, J.C., "Glycoproteins: what are the sugar chains for?" <u>TIBS</u> , 14:272-276 (1989)
C32	*Planas et al., "Reengineering the Catalytic Lysine of Aspartate Aminotransferase by Chemical Elaboration of a Genetically Introduced Cysteine," <u>Biochemistry</u> , 30:8268-8276 (1991)
C33	*Plettner, E., et al., "Modulation of Esterase and Amidase Activity of Subtilisin Bacillus Lentus by Chemical Modification of Cysteine Mutants," <u>Journal of the American Chemical Society</u> , (2 Jun. 1999) 121/21, 4977-4981, XPO000891274.
C34	*Plettner, Erika et al., "A Combination Approach to Chemical Modification of Subtilisin Bacillus Lentus," <u>Bioorganic & Medicinal Chemistry Letters</u> (Sept. 8, 1998) Vol. 8, No. 17, pp. 2291-2296, XP0004138220

EXAMINER DATE CONSIDERED

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy f this f rm with next communication to Applicant(s).

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FILING DATE	GROUP ART UNIT
July 30, 2003	Unassigned

OTHER DOCUMENTS (including Author, Title, Date, Pertinent Pages, Etc.) *Polgar et al., "A New Enzyme Containing a Synthetically Formed Active Site. Thiol-Subtilisin," C35 Journal of American Chemical Society, 88:3153-3154 (1966) Rademacher et al., "Glycobiology," C36 C37 *Ramachandran et al., "Stabilization of Barstar by Chemical Modification of the Buried Cysteines," Biochemistry, 35:8776-8785 (1996) C38 *Roberts et al., "Reactivity of Small Thiolate Anions and Cysteine-25 in Papain Toward Methyl Methanethiosulfonate," Biochemistry, 25:5595-5601 (1986) C39 *Siddiqui et al, "Arthrobacter D-Xylose Isomerase: Chemical Modification of Carboxy Groups and Protein Engineering Of pH Optimum," Biochem. J., 295:685-691 (1993) *Smith et al., "An Engineered Change in Substrate Specificity of Ribulosebisphosphate C40 Carboxylase/Oxygenase," The Journal of Biological Chemistry, 265:1243-1245 (1990) C41 *Smith et al., "Chemical Modification of Active Site Residues in y-Glutamyl Transpeptidase," The Journal of Biological Chemistry, 270:12476-12480 (1995) C42 *Smith et al., "Restoration of Activity to Catalytically Deficient Mutants of Ribulosebisphosphate Carboxylase/Oxygenase by Aminoethylation," The Journal of Biological Chemistry, 263:4921-4925 (1988)C43 *Smith et al., "Simple Alkanethiol Groups for Temporary Blocking of Sulfhydryl Groups of Enzymes," Biochemistry, 14:766-771 (1975) *Smith et al., "Subtle Alteration of the Active Site of Ribulose Bisphosphate Carboxylase/Oxygenase C44 by Concerted Site-Directed Mutagenesis and Chemical Modification," Biochemical and Biophysical Research Communications, 152:579-584 (1988) C45 *Spura, A., et al. "Probing Agonist Domain of the Nicotinic Acetylcholine Receptor by Cysteine Scanning Mutogenesis Reveals Residues in Proximity to the Alpha-Bungarotoxin Binding Site, Biochemistry, 20 Apr. 1999 Vol. 38:16 pp. 4912-4921 C46 *Stewart et al., "Catalytic Oxidation of Dithiols by a Semisynthetic Enzyme," J. Am. Chem. Soc., 108:3480-3483 (1986) C47 *Valenzuela et al., "Kinetic Properties of Succinylated and Ethylenediamine-Amidated δ-Chymotrypsins," Biochim. Biophys. Acta, 250:538-548 (1971) C48 *West et al., "Enzyme-catalysed Synthesis of Peptides Containing D-Amino Acids, J. Chem. Soc. Chem. Commun., pp 417-18 (1986) C49 *West et al., "Enzymes as Synthetic Catalysts: Mechanistic and Active-Site Considerations of Natural and Modified Chymotrypsin," J. Am. Chem. Soc., 112:5313-5320 (1990) C50 *White et al., "Sequential Site-Directed Mutagenesis and Chemical Modification to Convert the Active Site Arginine 292 Of Aspartate Aminotransferase to Homoarginine," Journal of the American Chemical Society, 114:292-293 (1992) C51 *Wynn et al., "Chemical Modification of Protein Thiols: Formation of Mixed Disulfides," Methods in Enzymology, 251:351-356 (1995)

EXAMINER DATE CONSIDERED

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FILING DATE July 30, 2003 GROUP ART UNIT Unassigned

OTHER DOCUMENTS (including Author, Title, Date, Pertinent Pages, Etc.)		
	C52	*Wynn et al., "Comparison of Straight Chain and Cyclic Unnatural Amino Acids Embedded in the Core of Staphylococcal Nuclease," <u>Protein Science</u> , 6:1621-1626 (1997)
	C53	*Wynn et al., "Mobile Unnatural Amino Acid Side Chains in the Core of Staphylococcal Nuclease," Protein Science, 5:1026-1031 (1996)
1	C54	*Wynn et al., "Unnatural Amino Acid Packing Mutants of Escherichia Coli Thioredoxin Produced by Combined Mutagenesis/Chemical Modification Techniques," Protein Science, 2:395-403 (1993)

EXAMINER

DATE CONSIDERED

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